Jeg. Alex

RESULT 5

AAX34988

AAX34988 standard; DNA; 30 BP.

AAX34988;

(first entry) 30-JUN-1999 Antisense oligonucleotide targeted to protein kinase A-RI-alpha gene.

Human protein kinase A-RI-alpha gene; antisense oligonucleotide;

colon cancer; lung cancer; stomach cancer; hepatic cancer; melanoma; carcinostatic; leukemia; large intestinal cancer; rectal cancer;

malignant lymphoma; tongue cancer; oesophagus cancer; breast cancer; uterus cancer; pharynx cancer; brain tumour; malignant myoma;

Synthetic.

Homo sapiens.

WO9616976-A1

X

.06-JUN-1996.

X

95WO-JP002452 01-DEC-1995; X 94JP-00324006. 02-DEC-1994;

(POKK ) POLA CHEM IND INC.

Tsuchiya M,

WPI; 1996-277711/28.

; 0 of the lung or stomach, hepatic cancer, malignant lymphoma, cancer of the tongue, oesophagus, breast, uterus or pharynx, brain tumour, melanoma, or leukaemia, large intestinal cancer, rectal cancer, colon cancer, cancer oligonucleotides is useful as a carcinostatic agent, e.g. for treating The present sequence represents an antisense oligonucleotide directed against the human protein kinase A-RI-alpha gene. The antisense . 0 Oligo:nucleotide contg. human protein kinase A gene sequence Length 30; Indels 0 U; 0 Other; .; 0 2; DB Pred. No. 7.5; 5; Mismatches Score 21; ω Claim 7; Page 16; 24pp; Japanese. % 8 GGCUGCGUGCCUCACUGG GGCTGCGTGCCTCCTCACTGG AAZ30811 standard; RNA; 30 BP. 100.08; 76.2%; 11 C; Conservative 3 A; carcinostatic agent. Best Local Similarity Sequence 30 BP; malignant myoma 05-JAN-2000 16; σ AAZ30811, Query Match AAZ30811/c Matches RESULT 6 ID PS XX Qy X  $\mathcal{C}$ 엄

Synthetic substrate RNA 30mer.

Oligonucleotide; phosphorothioate; phosphodiester; linkage; POPS block; stability; antisense; RNase H; activation; cleavage; gene therapy;

W immune effects; ss.

דווחוומוום פדדפרנאי

Synthetic.

WO9950409-A1

PD 07-0CT-1999

X

 $\stackrel{\times}{\sim}$ 

X

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X

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01-APR-1999; 99WO-US007276.

01-APR-1998; 98US-0080321P

PA (HYBR-) HYBRIDON INC.

Zhou WQ, Agrawal S;

WPI; 1999-610851/52.

specific phosphorothioate nucleosides, used to control the expression of Antisense oligonucleotides with alternating phosphodiester and genes.

PT

 $\approx$ 

Example 3; Page 11; 27pp; English.

(PO) and nucleoside phosphorothioates (PS), designated a POPS block. The This sequence represents a synthetic RNA 30mer, used as a substrate for comprise a region containing an alternating nucleoside phosphodiesters studies of the effects of novel antisense oligonucleotides (AAZ30809-Z30810) on RNase H activity. The novel antisense oligonucleotides olimning ontides have

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evaluate the effect of blocking such specific gene expression at selected
                                                                                                                                                                                                                                                                                   to control the expression of specific genes. They can be labelled with a
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         be used for therapy in the treatment of diseases resulting from aberrant
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 stages of development or differentiation. The oligonucleotides may also
                                                                                                                                                                                                                                         partial thromboplastin time. The antisense oligonucleotides can be used
                                                                                                                                                                                                      such as immune stimulation, complement activation and prolongation of
                                                                                                                                                                                                                                                                                                                                                                                                                                                    specific gene in an experimental cell culture or animal system and to
                                                                                                                                                           phosphorothioate oligonucleotide-related side effects of gene therapy
                                                                                                                                                                                                                                                                                                                                                                   hybridisation assays. They can also be used as antisense probes of
antisense oligonucleotides have a reduced phosphorothioate content
                                                                                stability, nuclease stability, RNase H activity, antisense-based
                                                                                                                                                                                                                                                                                                                                                                                                         specific gene function by being used to block the expression of
                                        without compromising their antisense properties, such as duplex
                                                                                                               biological activity and tissue disposition. They can reduce the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    .
0
                                                                                                                                                                                                                                                                                                                              reporter group and used as probes in conventional nucleic acid
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Indels
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               6 A; 10 C; 11 G; 0 T; 3 U; 0 Other;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Score 21; DB 2;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Pred. No. 7.5;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                5; Mismatches
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1 GGCUGCGUGCCUCCUCACUGG 21
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    gene expression (e.g., cancer)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        100.08;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             16;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Query Match
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BP.

ADP83666 standard; RNA;

RESULT 7 ADP83666

ΠD

ADP83666;

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disease caused by the overexpression or
                                                                        complement and reduced anti-thrombotic properties. By controlling the
                                                                                                                         proliferation of cancer cells and tumour growth is possible. This is
                                                                                                 regulation of protein kinase A subunit RI-alpha, inhibition of the
                                                  e.g. reduced mitogenicity, reduced activation of
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              inverted chimeric hybrid; protein kinase A subunit RI-alpha gene;
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       DNA/RNA hybrid; antisense; hybrid; inverted hybrid; mitogenicity,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Protein kinase A subunit RI-alpha synthetic oligonucleotide #165
                       oligonucleotides produced fewer side effects than conventional
                                                                                                                                                                                                                                                                                                                               ö
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           cell proliferation; tumour;
of protein kinase A subunit RI-alpha gene expression. Such
                                                                                                                                                                                                                                                                           DB 2; Length 18;
                                                                                                                                                                                                                                                                                                                               Indels
                                                                                                                                                                                                                            2 U; 0 Other;
                                                                                                                                                                                                                                                                                                 4.8e+02;
                                                                                                                                                                                                                                                                                                                             Mismatches
                                                                                                                                                                                                                                                                             Score 17;
                                                                                                                                                                                                                                                                                                   Pred. No.
                                                                                                                                                                          inappropriate expression of this gene
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                                                                                                                                                  method for the treatment of
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ribonucleotide; ss.

Synthetic.

Key Location/Qualifiers

misc\_RNA 1..4

1. .4
/\*tag= a
/note= "ribonucleotide"

/moce ribonacted the 15. 18 /\*tag= b

misc\_RNA

/note= "ribonucleotide"

WO9711171-A1.

27-MAR-1997.

X

19-SEP-1996; 96WO-US015084.

22-SEP-1995; 95US-00532979.

X

X

(HYBR-) HYBRIDON INC.

XX PI Agrawal S; WPI; 1997-202880/18.

- are useful for Modified protein kinase A specific oligo:nucleotide(s) the treatment of cancer,

Claim 4; Page 17; 66pp; English.

modified antisense oligonucleotide This oligonucleotide was found to (#165) which was designed as a hybrid. : + - - - + : This sequence represents a synthetic,

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0
                                                                                         gene
reduce tumour growth in mice if administered orally or by intraperitoneal
                                                                                                                                                                                                                                                           possible. This is a novel method for the treatment of disease caused by
                          injection. The modified oligonucleotide types used in this study were
                                                                                   investigate the down regulation of protein kinase A subunit RI-alpha
                                                     hybrid, inverted hybrid or inverted chimeric hybrid and were used to
                                                                                                                                                                                                                              inhibition of the proliferation of cancer cells and tumour growth is
                                                                                                                                                                           By
                                                                                                              expression. Such oligonucleotides produced fewer side effects than
                                                                                                                                          conventional oligonucleotides e.g. reduced mitogenicity, reduced
                                                                                                                                                                      activation of complement and reduced anti-thrombotic properties.
                                                                                                                                                                                                   controlling the regulation of protein kinase A subunit RI-alpha,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ;
0
                                                                                                                                                                                                                                                                                        the overexpression or inappropriate expression of this gene
                                                                                                                                                                                                                                                                                                                                                                                                         DB 2; Length 18;
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                                                                                                                                                                                                                                                                                                                                                  0 Other;
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                                                                                                                                                                                                                                                                                                                                                                                                                                  Best Local Similarity
                                                                                                                                                                                                                                                                                                                                                Sequence 18 BP;
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Protein kinase A subunit RI-alpha synthetic oligonucleotide #164

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ANTISENSE OLIGONUCLEOTIDES FOR TREATMENT OF CANCER
                                                                                                                                                              Sterne, Kessler, Goldstein and Fox
                                                                                                                                                                                 STREET: 1225 Connecticut Avenue, N.W.
                                                                                                                                                                                                                                                                                                                                                                                                                                        US/07/702,163B
Sequence 2, Application US/07702163B
                                                                                                                                                                                                                                                                                                                                               COMPUTER: IBM PC compatable
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       REGISTRATION NUMBER: 32,893
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    TELECOMMUNICATION INFORMATION:
                                                                      APPLICANT: Yoon S. Cho-Chung
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    none
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            TELEPHONE: (202) 466-0800
                                                                                                                                                                                                                                                                                                                          MEDIUM TYPE: Floppy disk
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ATTORNEY/AGENT INFORMATION:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               REFERENCE/DOCKET NUMBER:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 (202)833-8716
                                                                                                                                                                                                                                                                                                                                                                                                                CURRENT APPLICATION DATA:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Robert W. Esmond
                                                                                                                                                                                                                                                                                                                                                                                                                                                              19910520
                                                                                                                                                                                                                                                                                                   COMPUTER READABLE FORM:
                                                                                                                                         CORRESPONDENCE ADDRESS:
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    424
                                                                                                                                                                                                                                                                                                                                                                                                                                        APPLICATION NUMBER:
                                                                                                                                                                                                                                                                                                                                                                                             PatentIn
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 APPLICATION NUMBER:
                                                                                                                NUMBER OF SEQUENCES:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           none
                                                                                                                                                                                                                                                                                                                                                                     OPERATING SYSTEM:
                                                                                            TITLE OF INVENTION:
                                                                                                                                                                                                           Washington
                                                GENERAL INFORMATION:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    CLASSIFICATION:
                       Patent No. 5271941
                                                                                                                                                                                                                                                        COUNTRY: USA
                                                                                                                                                                                                                                                                                                                                                                                                                                                              FILING DATE:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       FILING DATE:
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С
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                                                                                                                                                                                                           CITY:
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US-07-702-163B-2

100 202010

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ANTISENSE OLIGONUCLEOTIDES FOR TREATMENT OF CANCER
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                                                                                                                                                                                         DB 2; Length 18;
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                                                                                                                                                                                                           Pred. No. 1.4e+02;
                                                                                                                                                                                                                               4; Mismatches
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                                                                                                                                                                                          Score 17;
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                Cho-Chung
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INFORMATION FOR SEQ ID NO:
                  SEQUENCE CHARACTERISTICS
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                                                        TYPE: NUCLEIC ACID
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                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Washington
                                     18 bases
                                                                                             TOPOLOGY: linear
                                                                                                               MOLECULE TYPE: DNA
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248636 SS1

TELEX: